

#16/A

Docket No.: 213.1079-CIP3

**UNITED STATES PATENT & TRADEMARK OFFICE**

Examiner: RUSSEL, J.E. Art Unit: 1654

Re: Application of: GREENWALD, R.B., et al.

Serial No.: 09/758,993

Filed: January 12, 2001

For: TETRAPARTATE PRODRUGS

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**AMENDMENT**

Assistant Commissioner  
for Patents  
Washington, D.C. 20231

March 7, 2003

Sir:

In response to the Office Action dated December 10, 2002 please amend the above—

identified application as follows:

**IN THE SPECIFICATION:**

**Please replace the paragraph beginning at page 2, line 20 with the following:**

One way in which these problems have been addressed is described, for example, by co-owned patent applications Serial Nos. 09/183,557, filed October 30, 1998, now U.S. Patent No. 6,180,095 and 08/992,435, filed on December 17, 1997, now abandoned. These teach double prodrugs, i.e., tripartate, that comprise polymer conjugates of various biologically-effective materials, and methods of making these conjugates. The double prodrug linkages are selected to hydrolyze *in vivo* at a rate which generates sufficient amounts of the “second” and more reactive prodrug compound within a suitable time after administration by, e.g., a 1,4-aryl or 1,6-aryl (e.g., benzyl) elimination reaction, providing improved control of the pharmacokinetics of a number of small molecule

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